REVIEW

Occupational risk of Lyme disease: an epidemiological review

J D Piacentino, B S Schwartz

Occup Environ Med 2002;59:75-84

Lyme disease is the most common vector borne disease in the United States. Since the early 1980s, a large body of literature has evaluated the occupational risk of Lyme disease. The availability of a new vaccine to prevent Lyme disease makes it necessary for occupational health professionals to make decisions regarding the occupational risk of the disease among employees.

method has been developed to categorise published studies into four groups based on their use in the assessment of the occupational risk of the disease (high, moderate, low, and none). This categorisation was based on study design, the definition of the occupational group, the presence and definition of the comparison group without occupational risk, the diagnostic basis for Lyme disease, and the method of laboratory confirmation. Four sources were used to obtain published articles (Medline, NIOSHTIC, Science Citation Index, and the European Union Concerted Action on Lyme Borreliosis website).

A total of 91 unique articles were reviewed for possible relevance to the occupational risk of Lyme disease, and 41 unique articles with primary data about occupational Lyme borreliosis were selected for detailed analysis. After applying the use for assessment method, 10 studies met criteria for high or moderate use; all but one were from European study populations.

Overall, the published literature suggested that outdoor workers may be at an increased risk of seropositivity for antibodies to *Borrellia burgdorferi*, mainly assessed in cross sectional studies with enzyme linked immunosorbent assay (ELISA) or indirect fluorescent antibody (IFA) only. However, many of these studies compared outdoor workers with groups that may have had lower risk of Lyme disease from residential and recreational exposure to ticks, limiting the inferences that can be made about occupational risk. Also, most of the studied seropositive workers did not have any symptoms compatible with Lyme disease, suggesting that the increased risk may be for asymptomatic infection.

Lyme disease is the most common vector borne illness in the United States. It is caused by infection with the bacterium *Borrelia burgdorferi* and transmitted by ticks of the *Ixodes ricinus* complex. Between 1992 and 1998, 88 967 cases of

Lyme disease were reported to the Center for Disease Control, with a mean annual incidence of 5.1 cases/100 000 population.2 During this period, there was a 70% increase in the annual number of reported cases, from 9909 in 1992 to 16 802 in 1998. This is likely to represent both a true increase in the incidence of the disease, and an increase in the proportion of diagnosed cases that were reported.2 However, it is estimated that for every reported case of Lyme disease, there are 7–12 cases that are unreported.^{3 4} Between 1992 and 1998, 10 states accounted for 92% of all reported cases of Lyme disease: New York (32.8%), Connecticut (17.4), Pennsylvania (14.6), New Jersey (12.2), Wisconsin (3.6), Rhode Island (3.5), Maryland (3.1), Massachusetts (2.4), Minnesota (1.7), and Delaware (1.0).2

The risk of Lyme disease would seem to be an important concern to outdoor workers in endemic areas. To date, Lyme disease has been documented in many occupational groups, including forestry workers, farmers, veterinarians, military recruits, orienteers, and outdoor workers in general. However, the risk of symptomatic infection in outdoor workers has not been well described and there are no recent systematic reviews of the topic. Published studies have mainly relied on measurement of the seroprevalence of antibodies to Borrelia burgdorferi among occupational groups compared with controls. Studies in the general population and outdoor workers have documented that the risk of the disease can decline over time, probably due to behavioural change and acquisition of immunity.5 6 Also, studies of outdoor workers have suggested that knowledge about Lyme disease, use of personal protective behaviour, and the development of pruritic reactions to tick bites may mitigate the risk of the disease among such workers.7-10 The result is an overall uncertainty about the potential occupational risk of Lyme disease in outdoor workers.

Before 1998, prevention of Lyme disease in outdoor workers was mainly limited to the use of personal preventive behaviour, including protective clothing, insect repellent, and tick checks and early tick removal.⁸ Environmental control methods are also available—such as control of grass and brush and application of insecticides. However, these methods have limited practicality for the prevention of Lyme disease in outdoor workers.¹¹ ¹² The

See end of article for authors' affiliations

Correspondence to: Dr B S Schwartz, Division of Occupational and Environmental Health, Johns Hopkins Bloomberg School of Hygiene and Public Health, Room 7041, 615 North Wolfe Street, Baltimore, MD 21205, IISA

Accepted 27 July 2001

Abbreviations: EUCALB, European Union Concerted Action on Lyme Borreliosis; ELISA, enzyme linked immunosorbent assay; IFA, indirect fluorescent antibody; PCR, polymerase chain reaction

Table 1 Distribution of published articles by geographical location and author among 41 articles with primary data on the occupational epidemiology of Lyme disease

Country	n (%)*	Author
Argentina	1 (2.4)	Stanchi and Balague 1993 ¹⁷
Austria	1 (2.4)	Schmutzhard et al 1988 ¹⁸
Croatia	1 (2.4)	Golubic et al 199819
Finland	1 (2.4)	Oksi and Viljanen 1995 ²⁰
France	2 (4.9)	Christiann et al 1996 ²¹ †; Zhioua et al 1997 ²²
Germany	2 (4.9)	Hauser et al 1998 ²³ ; Rath et al 1996 ²⁴
Ireland '	1 (2.4)	Robertson et al 1998 ²⁵
Italy	2 (4.9)	Nuti et al 1993 ²⁶ ; Santino et al 1998 ²⁷
Japan	2 (4.9)	Ikushima et al 1999 ²⁸ ; Nakama et al 1994 ²⁹
Lithuania	1 (2.4)	Montejunas et al 1994 ³⁰
Netherlands	5 (12.2)	Kuiper et al 1993 ³¹ ; Kuiper et al 1991 ³² ; van Charante et al 1998 ³³ ; van Charante et al 1994 ³⁴ ; Vos et al 1994 ³⁵
Poland	1 (2.4)	Chmielewska-Badora 1998 ³⁶
Spain	2 (4.9)	Arteaga et al 1998 ³⁷ ; Oteo et al 1992 ³⁸
Sweden	1 (2.4)	Gustafson et al 1993 ³⁹
Switzerland	2 (4.9)	Fahrer et al 1998 ⁴⁰ ; Fahrer et al 1991 ⁴¹
United Kingdom	6 (14.6)	Baird et al 1989 ⁴² ; Gregory et al 1993 ⁴³ ; Guy et al 1989 ⁴⁴ ; Morgan et al 1989 ⁴⁵ ; Reese and Axford 1994 ⁴⁶ ;
3		Thomas et al 1998 ⁴⁷
United States of America	10 (24.4)	Bowen et al 1984 ⁴⁸ ; Goldstein et al 1990 ⁴⁹ ; Klein 1995 ⁵⁰ ; Lane et al 1992 ⁵¹ ; Ley et al 1995 ⁵² ; Parrott et al 1993
	. ,	Schwartz and Goldstein 1990°; Schwartz et al 1993 ⁵³ ; Schwartz et al 1994 ⁵⁴ ; Śmith et al 1988 ⁵⁵
Total	41 (100)*	

*May not total 100% due to rounding; †control data supplied by Christiann et al 1997. 57

approval of a Lyme disease vaccine by the Food and Drug Administration offers a new method of prevention, independent of current strategies. Despite proved safety and 90% efficacy after three doses in subjects under 65 years of age, the role of the vaccine has been the focus of considerable discussion. 1 13-16

The Center for Disease Control has recommended that the vaccine be considered for "persons who reside, work, or play in areas of high or moderate risk". Specifically, "Lyme disease vaccination should be considered for persons aged 15-70 years who engage in activities (for example, recreational, property maintenance, occupational, or leisure) that result in frequent or prolonged exposure to tick-infested habitat". The current recommendation that the vaccine be considered for use in selected people at moderate to high risk, and an incomplete understanding of the occupational risk of Lyme disease, have led to indecision on the part of health care professionals, government agencies, and employers on the role of vaccination against Lyme disease in outdoor workers. In an effort to evaluate the need for Lyme vaccination in outdoor workers, we assessed the occupational risk of Lyme disease in the published, scientific literature.

METHODS

Identification and selection of published articles

An attempt was made to collect every article published in English in the scientific literature that contained any primary data on occupational risk factors for Lyme borreliosis. The following databases were searched: Medline, NIOSHTIC (NI-OSHTIC was produced by the National Institute of Occupational Safety and Health (NIOSH) and was distributed as OSH-ROM; the July 1998 version, from Silverplatter Information, Norwood, MA, was used), Science Citation Index, and the EUCALB (European Union Concerted Action on Lyme Borreliosis) website. A general search of Medline in October 1999 conducted on "keyword = Lyme disease" found 4917 published articles. In an effort to select only those articles with occupational relevance, "keyword = Lyme disease" was then combined with multiple other keywords including: agriculture (n=3 articles), work(er) (n=67), employee (n=4), outdoor (n=28), occupation(al exposure) (n=9), park (n=58), exposure (n=187), military (n=18), vaccine (n=220), forest(er) (n=66) and farm(er) (n=11). A general search of NIOSHTIC for "Lyme" also yielded 25 articles for potential review. Additionally, all of the articles referenced on the EUCALB website as of January 2000 (n=260) were assessed for occupational relevance. Finally, the Science Citation Index was used to cross reference any article that had cited any of the following authors in its references: Arteaga F, Christiann F, Fahrer H, Gustafson R, Guy E, Kuiper H, Nuti M, Rath P, Santino I, Schwartz B, and Zhioua E. These authors were selected because they were most often cited in articles of occupational risk of Lyme disease.

Each database yielded multiple articles, many of which were cross referenced between and within databases. Through these techniques, 91 unique articles were preliminarily reviewed for possible relevance to the occupational risk of Lyme disease. Only articles which contained primary data on occupational populations were considered for further analysis. Data were considered primary if they were collected by the researchers and referenced to an occupational population or exposure. The 50 rejected papers made no connection between the study data and an occupational population or exposure. Ultimately, 41 unique articles with primary data on occupational Lyme borreliosis were selected for detailed analysis, representing a broad range of geographic locations and 36 unique authors (table 1). 7 9 17-55 One additional article by Zhioua et al,56 was identified but was not reviewed separately because the data were derived from another included article (Fahrer et al).41 Control data for Christiann et al were obtained from a subsequent publication by Christiann et al.21 5

Evaluation of use of articles

Initial criteria were established to categorise articles according to their use in assessing the occupational risk of Lyme disease (table 2). In developing these criteria, the primary motivation was the recent approval of a vaccine to prevent Lyme disease by the Food and Drug Administration and its impact on the occupational health care environment. Current Occupational Safety and Health Administration requirements for vaccination of workers are limited to vaccination for hepatitis B virus in its bloodborne pathogens standard (29 CFR Part 1910.1030). This standard is designed to protect workers from a symptomatic, often serious, and sometimes fatal, illness. In evaluating the occupational risk of Lyme disease, similar attention was given to the morbidity associated with Lyme disease. Risk of symptomatic, clinically and laboratory confirmed infection was thus the primary outcome of interest.

Five major factors were evaluated in defining the use of the published articles for assessing the occupational risk of Lyme disease (table 2):

	Criteria used to stratify articles according to their use in assessment of
occupation	onal risk of Lyme disease

Use	Criteria	
High	Design	Prospective study design Clearly defined occupational group Clearly defined comparison group without occupational risk
	Diagnosis	Clinical diagnosis of Lyme borreliosis (generally by physician) Laboratory confirmation (by western blot*)
Moderate	Design	Cross sectional or case-control study design Clearly defined occupational group Clearly defined comparison group without occupational risk
	Diagnosis	Clinical diagnosis of Lyme borreliosis (generally by physician) Positive ELISA or IFA serology
Low	Design	Any design Clearly defined occupational group Clearly defined comparison group without occupational risk
	Diagnosis	Diagnosis based on self reported symptoms or serology only Positive ELISA or IFA serology
None	Design	Any design No defined occupational group No defined comparison group without occupational risk
	Diagnosis	Diagnosis based on self report Positive ELISA or IFA serology

^{*}Although other methods of laboratory confirmation were acceptable (culture, polymerase chain reaction), no studies used any method more definitive than western blot.

- (1) Study design prospective studies had higher utility than did case-control or cross-sectional studies;
- (2) The occupational group had to be clearly defined;
- (3) A comparison group without occupational risk had to be included;
- (4) Definition of Lyme disease, with symptomatic, clinically confirmed infection of primary interest, by contrast with definition of cases on the basis of self reported symptoms ascertained by questionnaire or asymptomatic seropositivity only;
- (5) Laboratory confirmation had to be included, with western blot and the definition of the Centers for Disease Control considered to be better documentation than enzyme linked immunosorbent assay (ELISA) or indirect fluorescent antibody (IFA) only.⁵⁸ No previous studies of the occupational epidemiology of Lyme disease used more rigorous definitions of

laboratory confirmation—such as culture or polymerase chain reaction (PCR) of skin tissue from *erythema migrans* lesions.

Measurement of occupational risk

After application of the criteria of usefulness to the 41 published articles with primary data on the occupational epidemiology of Lyme disease, two were defined as of high use, eight as moderate, 14 as low, and 17 had no use (table 3). For the 10 studies in the high or moderate categories, three epidemiological effect measures are reported and calculated, when possible, from the published data. These were, in order of least useful to most useful measure:

(1) Seroprevalence (odds) ratio—the prevalence of positive serological test results in the defined occupational group under study compared with the prevalence in a defined control group without occupational risk;

Use	n (%*)	Study design	Author
High	2 (4.9)	Prospective	Kuiper et al 1993 ³¹ ; Vos et al 1994 ³⁵
Moderate	8 (19.5)	Case-control Cross sectional Prospective	Hauser et al 1998 ²³ Chmielewska-Badora 1998 ³⁶ ; Gustafson et al 1993 ³⁹ ; Kuiper et al 1991 ³² ; Zhioua et al 1997 ²² Bowen et al 1984 ⁴⁸ †; Fahrer et al 1998 ⁴⁰ ‡; Fahrer et al 1991 ⁴¹
Low	14 (34.1)	Case-control Cross sectional	Stanchi and Balague 1993 ¹⁷ ; Christiann <i>et al</i> 1996 ²¹ § Arteaga <i>et al</i> 1998 ³⁷ ; Baird <i>et al</i> 1989 ⁴² ; Reese and Axford 1994 ⁴⁶ ; Robertson <i>et al</i> 1998 ²⁵ ; Smith <i>et al</i> 1988 ³⁵ ; Goldstein <i>et al</i> 1990 ⁴⁹ ; Schwartz <i>et al</i> 1993 ⁵³ ; Schwartz and Goldstein 1990 ⁹
		Prospective	van Charante et al 1998 ³³ ; van Charante et al 1994 ³⁴ ; Oksi and Viljanen 1995 ²⁰ ; Rath et al 1996 ²⁴
None	17 (41.5)	Case-control Case series Cross sectional	Ley et al 1995 ⁵² Gregory et al 1993 ⁴³ ; Golubic et al 1998 ¹⁹ ; Klein 1995 ⁵⁰ ; Guy et al 1989 ⁴⁴ ; Ikushima et al 1999 ²⁸ ; Morgan et al 1989 ⁴⁵ ; Nakama et al 1994 ²⁹ ; Nuti et al 1993 ²⁶ ;
		Prospective	Oteo et al 1992 ³⁸ ; Santino et al 1998 ²⁷ ; Schmutzhard et al 1988 ²⁸ ; Lane et al 1992 ⁵¹ ; Montejunas et al 1994 ³⁰ ; Parrott et al 1993 ⁷ ; Schwartz et al 1994 ⁵⁴ ; Thomas et al 1998

Reference (first author)	Study design, dates	Location	Study population v comparison group	Definition of Lyme disease or seroprevalence	Results of relevance to occupational risk
Arteaga 1998 ³⁷	Cross sectional, 1998†	Vizcaya, Spain	302 Outdoor workers (117 forestry, 52 large animal vets, 18 shepherds, 27 apiculturists, 74 mushroom and truffle	ELISA, WB (CDC criteria for serological reactivity), and self reported clinical	Seroprevalence outdoor workers = 15% (44/302) by WB
			gatherers, 14 other) v none	questionnaire.	Cumulative clinical prevalence, outdoor workers = 15% (11/44) by WB
Baird 1989 ⁴²	Cross sectional, 1989†	Wigtownshire, UK	101Samples from farmers, foresters, and gamekeepers and seven samples from patients with potential LD v none	IFA, ELISA and physician diagnosis	Seroprevalence = 11% (12/108) Cumulative clinical prevalence = 91% (11/12)
Bowen 1984 ⁴⁸	Prospective, 1978-82	Monmouth County, NJ, USA	366 Outdoor workers at naval weapons station v 766 indoor workers at naval weapons station	IFA, clinical interview and medical record review	Incidence outdoor workers = 3.8% (14/366)
Chmielewska-	Cross sectional, 1998†		1153 Workers exposed to ticks (880 forestry workers and		Incidence indoor workers = 0.8% (6/766), p<0.001 Seroprevalence farmers = 38.6%
Badora 1998 ³⁶	0.000 000.10.1.1, 1770	202111, i oldila	273 farmers), 458 patients suspected of LD (362 from neurological clinic and 96 from dermatologic clinic) v 100 healthy blood donors	in, y east, and physician diagnosis	Seroprevalence foresters = 28.1% Seroprevalence blood donors = 6%, p<0.001
			,		Cumulative clinical prevalence farmers/foresters = 0.0% (1/1153)
Christiann 1996 ²¹	Case-control with control data	Berry, France	59 Cases of Lyme disease among the residents of Berry Sud (Christiann 1996); 170 recreational hunters v 182	IFA or ELISA until Nov 1993 and then ELISA only; clinical examination and	58% (34/59) Of LD cases were among farmers
and Christiann 1997 ⁵⁷	supplemented from		blood donors (Christiann 1997)	WHO recommendations 1993 and	Seroprevalence hunters = 15% (25/170) Relative risk of seroreactivity hunters v blood donors = 1.79, p=0.00001
Fahrer 1998 ⁴⁰	Christiann 1997 Phase II, prospective,	Switzerland	Phase II, 305 seropositive orienteers reexamined v phase I,	CDC criteria 1990 FUSA self-reported clinical	Annual clinical incidence phase II orienteers = 0.8%
ramer 1770	1986–88 and 1993	OWIIZCIIGIIG	950 orienteers	questionnaire, physician diagnosis and	6 Month clinical incidence phase I orienteers = 0.8%
				medical record review.	Cumulative clinical prevalence phase II orienteers = 4.9% (15/305)
Fahrer 1991 ⁴¹	Phase I of Fahrer 1998, prospective,	Switzerland	950 Swiss orienteers v 51 healthy volunteers who had spent most of their life at altitudes >1000 m and 50	ELISA, self reported questionnaire, physician diagnosis and medical record	Cumulative clinical prevalence orienteers = 1.9%–3.1% Seroprevalence orienteers = 26.1%
	1986		inhabitants of Berne at altitude 500 m‡	review.	Seroprevalence high altitude = 3.9% (2/51)
					Seroprevalence Berne = 6.0% (3/50)
					Six month clinical incidence orienteers = 0.8%
Goldstein 1990 ⁴⁹	Cross sectional, Oct 1998	New Jersey, USA	689 Employees from the NJ Natural and Historic Resources Section from 12 different sites v none	IFA, ELISA, and self reported clinical questionnaire	Seroprevalence = 5.7% (39/689) by IFA or ELISA There was no association between seropositivity and job title or job habitat
Golubic 1998 ¹⁹	Case series, 1998†	Croatia	218 Cases of LB in NW Croatia v none	IFA, self reported questionnaire, physician diagnosis and medical record review.	Clinical prevalence according to occupation: student 15% (33/218), agricultural worker 30% (30/218), agricultural clerk 14% (32/218), pensioner 12.5% (27/218) and housewife 14% (30/218)
Gregory 1993 ⁴³	Case series, 1987–91	UK	Two cases of neuroborreliosis and 6 more reports of clinical LD with positive ELISA v none	ELISA and physician diagnosis	2 Case reports of neuroborelliosis and six case reports of LD among military personnel in the U
Gustafson 1993 ³⁹	Cross sectional, Oct 1990	Stockholm, Sweden	362 Orienteers from the county of Stockholm during a large relay race in October 1990	ELISA, self reported clinical questionnaire and physician diagnosis	Seroprevalence orienteers = 9% (31/362) Seroprevalence controls = 2% (A-1/50), 9% (B-13/150), 1% (C-1/74), 2% (D-9/378)
			A - 50 Blood donors		Seroprevalence OR orienteers v control B = 0.9 (95% CI 0.5 to 1.8)
			B - 150 People living in Sweden (no orienteers)		Seroprevalence OR orienteers v control C = 9.3 (95% Cl 3.6 to 24.2)
			C - 74 Hospital patients D - 378 People from Iceland (no Ixodes ticks)		Cumulative clinical prevalence orienteers = 6% (22/362) Cumulative clinical prevalence controls-B = 1% (2/150), OR = 4.8 (95% CI 1.1 to 20.6)
Guy 1989 ⁴⁴	Cross sectional, 1989†	Southampton, UK	41 Forestry Commission workers (11 keepers, 30 other) v	ELISA, WB and self reported clinical	Seroprevalence = 25% (10/40) by WB
•		•	none	questionnaire	Cumulative clinical prevalence = 5% (2/40)
Hauser 1998 ²³	Case control, 1989–95	Germany	222 Patients with clinically defined LB and 458 asymptomatic forestry workers v 133 blood donors	ELISA and physician diagnosis	Seroprevalence foresters = 41–44.8%, depending on ELISA antigen.
Ikushima 1999 ²⁸	Cross sectional, 1999†	lanan	80 Forestry workers v none	ELISA and WB	Seroprevalence blood donors = 8%, OR = 8.4 (95% CI 4.4 to 15.9) Seroprevalence foresters = 22.5% (18/80) by WB
Klein 1995 ⁵⁰	Case series, 1995	Wilmington, DE,	Five physicians amona 83 employed pediatricians and 55		One year clinical prevalence physicians = 3.6% (5/138)
Kuiper 1993 ³²	Prospective, 1989–90	USA Netherlands	pediatric residents v none 151 Dutch forestry workers v 151 male office workers	ELISA, WB, and physician diagnosis with reference to CDC case definition,	Seroprevalence forestry = 28% (43/151) by ELISA
			matched for age and residence	1990	Seroprevalence office = 5% (8/151) by ELISA, p<0.01 Clinical incidence forestry = 0.0%
					Seroconversion forestry = 5% by ELISA/WB
					18% (7/39) Of seropositive forestry workers or 5.5% (7/127) of all forestry workers in 1989 the case definition criteria for LB.
Kuiper 1991 ³¹	Cross sectional, 1989	Netherlands	127 Dutch forestry workers v 127 male office workers,	IFA, WB and physician diagnosis	Seroprevalence foresters = 19.7% (25/127)
			matched for age and region	(adapted CDC classification)	Seroprevalence office = 6.3% (8/127), OR = 3.7, 95% CI 1.5 to 9.7.
Lane 1992 ⁵¹	Prospective, 1988-89	Northwest	119 Residents of Mendocino County at entry (99 current	IFA, WB and physician diagnosis	Cumulative clinical prevalence foresters = 6% (7/127) Seropositivity was associated with greater years of residence in the area (p=0.032), decreased
1772		California	and 20 former residents) and 59 at follow up v none	, and physician diagnosis	hiking (p=0.006), and woodcutting (p=0.048).
100552	6	C lif	All (FAA . I. CAD CH III	N · · · · ·	Time spent working outdoors was not identified as a risk factor for probable LD.
Ley 1995 ⁵²	Case control, June 1991–Dec 1992	California	All cases of EM reported to CA Department of Health Services v age and sex matched controls	Physician diagnosis	Diagnosis of LD was not associated with work outdoors, OR = 1.04 (p=0.87) or total number of hours spent outside during leisure activities per month.
Montejunas 1994 ³⁰	Prospective, 1988-91	Lithuania	Three occupational groups: foresters (n=268), outside field	IFA	Seroprevalence foresters = 14% (37/268)
1994			Three occupational groups: foresters (n=268), outside field workers (n=115), and veterinarians (n=68) v 163 urban industrial workers		Seroprevalence field = 22% (25/115)
					Seroprevalence vets = 32% (22/68) Seroprevalence urban = 4% (6/163), OR = 2.5, p<0.001

Reference (first author)	Study design, dates	Location	Study population v comparison group	Definition of Lyme disease or seroprevalence	Results of relevance to occupational risk
Morgan 1989 ⁴⁵	Cross sectional, 1989	UK	180 Farmers and their familes v 75 control patients who lived in the area, but who denied contact with farm	ELISA	Seroprevalence farmers/families = 14.4% (26/180)
			animals		Seroprevalence control = 2.6% (2/75)
Nakama 1994 ²⁹	Cross sectional, 1990–91	Nagano, Japan	222 Forestry workers v 760 residents of an agricultural town	IFA	Seroprevalence foresters = 1.1% [8/760] Seroprevalence residents = 5.9% [13/222], p<0.01
Nuti 1993 ²⁶	Cross sectional, 1987–91	Italy	1146 Subjects subdivided into six categories: farmers (395), forestry workers (265), rangers (82), soldiers (299), hunters (75) and fishermen (30) v none	IFA	Seroprevalence farmers = 10.1% (40/395) Seroprevalence foresters = 19.6% (52/265) Seroprevalence rangers = 19.5% (16/82) Seroprevalence soldiers = 3.0% (9/299) Seroprevalence hunters = 8.0% (6/75)
Oksi 1995 ²⁰	Prospective, Jun 1993–Dec 1993	Gylto, Finland	77 Military recruits in Lyme endemic area initially, 67 recruits completed the study at 6 months v 50 military recruits in nonendemic area, initially; 33 recruits completed the study at 6 months.	ELISA and self reported clinical questionnaire	Seroprevalence fisherman = 16.6% (5/30) No probable history of EM in either group. Seroprevalence military endemic = 16.9% (13/77) Seroprevalence military nonendemic = 4.0% (2/50) No change in IgG seroprevalence at 6 month follow up for either group.
Oteo 1992 ³⁸	Cross sectional, Oct 1986–Mar 1988	La Rioja, Spain	500 Non-randomised individuals residing in Rioja, Spain v none	IFA and self reported clinical questionnaire	Seroprevalence outdoor worker = 20% v other 4.7%, p<0.001 Seroprevalence was associated with rural residence, p<0.001 Seroprevalence was associated with foresters, cattle raisers, and contact with domestic animals, p<0.001
Parrott 1993 ⁷	Prospective, May 1989–Oct 1989	Assateague Island, MD	99 Outdoor workers on Assateague Island initially with 86 workers continuing to post seasonal evaluation v None	EUSA and self reported clinical questionnaire	28% of farmers/foresters showed clinical signs compatible with LD Seroprevalence = 0% Seroconversion 3 month follow up = 0%
Rath 1996 ²⁴	Prospective, Feb-Sep 1992	Brandenburg, Germany	626 Foresters initially, 406 foresters at 6 month follow up v 200 blood donor controls	IFA, IBA, and self reported clinical questionnaire	Clinical prevalence = 0% Seroprevalence foresters = 8% by IBA Seroprevalence controls = 4% by IBA, p<0.05 Seroconversion foresters, 6 month follow up = 7.2% (IBA)
Reese 1994 ⁴⁶	Cross sectional, 1993	London, UK	44 Outdoor park workers from Richmond and Bushey parks v 27 zoo keepers from Whipsnade wildlife park in Bedfordshire, who worked in a similar outdoor	ELISA, IBA and self reported clinical questionnaire	Seroprevalence park = 32% (14/42) by IBA (three bands) Seroprevalence zoo = 4% (1/27) by IBA (three bands), p<0.005
Robertson 1998 ²⁵	Cross sectional, 1998†	Ireland	environment 38 National park rangers v 1224 blood donors from the same location as the park rangers	ELISA and IBA (5 bands present)	Seroprevalence park rangers = 0% (0/22) by IBA Seroprevalence blood donors = 3.4% (42/1224) by IBA
Santino 1998 ²⁷	Cross sectional, Jun–Aug 1995	Abruzzo, Italy	22 Park workers at an altitude of 750 to 1150 m v 50 park inhabitants at an altitude of 1150 m‡	ELISA, WB, and self reported clinical questionnaire	Seroprevalence blood admors = 9.1% (42/21) by ELISA, altitude = 800 m, 1080 m Seroprevalence park rangers = 4.5% (1/22) by WB (5 bands), altitude unknown Seroprevalence park inhabitants = 0.0% by ELISA, altitude = 1150 m None of the park workers or inhabitants showed signs compatible with LD.
Schmutzhard 1988 ¹⁸	Cross sectional, 1985	Tyrol, Austria	80 Austrian Federal Army soldiers intially, 50 soldiers at 4 week follow up (serology) and clinical observation for 14 weeks ν none	ELISA and physician diagnosis	Seroprevalence at 4 week follow up = 38% (18/50) Seroconversion = 22% (11/50) Clinical prevalence at 14 week follow up = 4% (2/50)
Schwartz 1990 ⁹	Cross sectional, Sep–Oct 1988	NJ, USA	689 Employees of NJ State Dept. of Env. Protection, included both indoor and outdoor workers <i>v</i> subset of indoor workers	IFA, EUSA, and self reported clinical questionnaire	Critical prevalence in 14 week follow up = 4% (2/30) Seroprevalence indoor/outdoor workers = 5.7% (39/689) Crude OR associated with occupational tick exposure = 2.2 (95% CI 0.7 to 9.0). Adjusted OR associated with occupational tick exposure = 5.1 (95% CI =1.1 to 23.6).
Schwartz 1993 ⁵³	Cross sectional, Oct 1990	NJ, USA	758 Employees of NJ State Dept. of Env. Protection Outdoor workers <i>v</i> none	ELISA and self reported clinical questionnaire	Seroprevalence outdoor, 1988 = 8.1% Seroprevalence outdoor, 1990 = 18.7% (142/758) LD incidence in the general population increased 30% from 1989 to 1991
Schwartz 1994 ⁵⁴	Prospective, 1988–91	NJ, USA	NJ outdoor workers from Dept. of Env. Protection; 1,519 workers for at least 1 y, 378 workers for 2 y, 228 for 3 yrs, and 192 for 4 years v none	IFA in 1988 and then ELISA from 1989 to 1991; and self reported clinical questionnaire	Seroprevalence outdoor = 4.4 to 18.7% Seroconversion outdoor = 0.6 to 16.7% Seroreversion outdoor = 23 to 53% Risk factors for seroconversion included years at residence, rural residence, pet ownership, and history of medical problems.
Smith 1988 ⁵⁵	Cross sectional, May–Nov 1986	NY, USA	414 State employees of the NY State Office of Parks, Recreation and Historic Preservation and of the NY State Department of Environmental Conservation v 362 NY State blood donors in Lyme endemic area and NY State blood donors in non Lyme endemic area, total number unknown	EUSA, WB and self reported clinical questionnaire	Seroprevalence state employees = 6.5% (27/414) Seroprevalence Lyme endemic controls = 1.1% (4/362) Seropositivity RR state employees v Lyme endemic controls = 5.9 (95% CI 2.4 to 14.6) RR seropositivity outdoor worker v indoor worker = 2.0 (95% CI 0.3 to 13.0). Seropositivity was associated with leisure time outdoor exposure, while the evidence for an association with work exposure was less consistent. Hours spent outdoors during work was not associated with seropositivity.

Table 4 continued	ontinued				
Reference (first author)	Study design, dates Location	Location	Study population v comparison group	Definition of Lyme disease or seroprevalence	Results of relevance to occupational risk
Stanchi 1993 ¹⁷	Stanchi 1993 ¹⁷ Case control, 1993 Argentina	Argentina	28 Farmers referred by physicians for arthritis v 25 farmers without past or present arthritis	IFA	Seroprevalence farmers with arthritis = 11% (3/28) Seroprevalence farmers without arthritis = 0% (0/25)
Thomas 1998 ⁴⁷	Prospective, 1991 to UK 96	UK	404 Farmers and their families at enrollment; 387 at 1 year, 345 at 2 years, and 336 at 3 years v None	ELISA, WB and self reported clinical questionnaire	Seroprevolence enrollment = 0.2% (1/404) by WB Seroreversion = 0.3% (1/387) Seroconversions = None
Van Charante 1998, ³³ Follow up of Van Charante 1994 ³⁴	Prospective, 1989 to Netherlands 93	Netherlands	312 Forestry workers and muskrat catchers straified according to high or low exposure v 356 office workers with no exposure	ELISA, WB self reported clinical questionnaire	Seroprevalence OR high exposure foresters/muskrat v office workers = 17 (95% CI 6.4 to 5.5) Seroprevalence OR low exposure foresters/muskrat v office workers = 6.1 (95% CI 2.9 to 13) Seroconversion = 0.23 vear - (95% CI 0.12 to 0.34)
Van Charante 1994 ³⁴	Prospective, 1989 to Netherlands 90	Netherlands	151 Forestry workers v 151 office clerks	ELISA, self reported clinical questionnaire	Seroprevelence forester 19.9% (30/151) Seroprevelence office = $6\% (9/151)$ Seroprevelence office = $6\% (9/151)$ Seroprevelence OR foresters v office = $3.9 (95\% CI 1.7 to 9.7)$
Vos 1994 ³⁵	Prospective, Jan to Dec 1991	Netherlands	Initially, 905 outdoor military recruits. After 1 year ELISA, WB 469 recruits for second blood draw v Initially, 1253 questionna indoor military recruits. After 1 year 463 for follow up diagnosis	, self reported clinical ire and physician	Seroconversion indoor = 0.9% $(4/469)$ Seroconversion outdoor = 2.2% $(10/463)$ Seroconversion RR outdoor v indoor = 0.4 $(95\% \text{ Cl} \ 0.1 \text{ to} \ 1.2)$ Clinical incidence among seroconverted outdoor = 7.1% $(1/1/4)$
Zhioua 1997 ²²	Cross sectional, 1997	France	212 Forestry workers v 31 blood donors with no contact with the forest	IFA and physician diagnosis	James increases a month g succentration index g
*Abbreviations c density; OR, odc	as in table 1; CDC, Cent ds ratio; RR, relative risk;	ters for Disease Cor WB, western blot;	ntol; EUSA, enzyme linked immunosorbent assay; EM, er, Tindicate date of publication, no information provided re	thema migrans; IBA, immunoblot assay garding dates for collection of data; ‡al	*Abbreviations as in table 1; CDC, Centers for Disease Control; EUSA, enzyme linked immunosorbent assay; EM, erythema migrans; IBA, immunoblot assay; IFA, immunoflourescent antibodies; LB, Lyme borreliosis; LD, Lyme disease; OD, optical density; OR, odds ratio; RB, western blot; Tindicate date of publication, no information provided regarding dates for collection of data; taltitudes > 1000 m represent non-endemic areas for Lyme disease.

(2) Clinical prevalence (odds) ratio—the prevalence of symptomatic, clinically confirmed Lyme disease with laboratory confirmation in the defined occupational group compared with the prevalence in the control group;

(3) Clinical incidence ratio—the incidence of symptomatic, clinically confirmed Lyme disease with laboratory confirmation in the defined occupational group compared with the incidence in the control group. When calculated by the author of the published study, these data were reported directly. When sufficient data were included in the article to calculate a desired epidemiological effect measure, but not reported by the author, the effect measure was calculated using Intercooled Stata™, version 6 (Stata Corporation, College Station, TX)

RESULTS

Descriptive summary of published literature

The published literature on the occupational epidemiology of Lyme disease included 41 articles with a mean (SD) occupational group sample size of 367 (378) and a mean (SD) comparison group sample size of 196 (333) (table 4). Of these studies, two (4.9%) were conducted (or initiated) before 1985; 23 (56.1%) were conducted from 1986 to 1990; 10 (24.3%) were conducted from 1991 to 1995; and six (14.6%) were conducted from 1996 to 1999. A total of 14 (34.2%) used a prospective study design (including one study with a retrospective cohort design, by Bowen *et al* (1984)⁴⁸); 20 (48.8%) were cross sectional in design; four (9.8%) were case-control studies; and three (7.3%) were case series. Sixteen (39%) studies presented data only for occupationally exposed subjects without reference to a control group, and 25 studies (61%) presented data for both occupational and comparison groups.

The definition of Lyme disease varied among the studies. Twenty three (56.1%) studies solely relied on serological assessment. Sixteen (39.0%) studies used both serological assessment and clinical evaluation by a physician and two (4.9%) studies used only clinical evaluation by a physician without any laboratory confirmation. Of the 39 studies with serological data, 15 (38.5%) confirmed positive results by ELISA or IFA with western blot.

Evaluation of occupational risk in studies of high and moderate use

Of the 10 studies with high and moderate use, seven reported the prevalence of seropositivity among the occupational group under study, which ranged from 1.0% to 44.8%. When compared with the seroprevalence among controls, six of the seven studies documented a significantly increased relative odds of seropositivity ranging from 3.7 to 9.9 (table 5), whereas the association found in one study (Zhioua *et al*²²) did not reach statistical significance. However, only two of the seven studies (Kuiper et al^{31 32}) matched the comparison subjects to the occupationally exposed subjects for age and residence. The other studies did not reach an important goal of the comparison group, which would be the ability to separate occupational risk from residential and recreational risk, because it seemed that comparison subjects also had a lower risk of residential and recreational tick exposure. Of these seven studies, only two studies (Kuiper et al31 32) performed western blots as confirmatory evidence in subjects with positive ELISA or IFA test results according to current standards recommended by the Centers for Disease Control.⁵⁸ It should be noted that all seven studies were performed in Europe where confirmatory western blots are not standardised and four of the seven studies predated the 1995 Centers for Disease Control criteria for Lyme disease serological testing.

Of the 10 studies with high and moderate use, six presented data on the prevalence of clinically confirmed Lyme disease in the occupational group under study, ranging from 0.4% to

	Seroprevalence			Clinical prevalence	lence		Clinical incidence	nce	
Study	Study group (%)	Control group (%)	OR (95% CI)	Cor Study group (%) (%)	Control group %) (%)	OR (95% CI)	Study group (%)	Control group (%)	RR (95% CI)
High use studies (n=2)	C	0	+10 21 -10 67 1 7	, E	2	<u> </u>		2	\frac{\lambda}{2}
Kuiper et al 1993	87	0.0	7.1 (3.2 to 13.8)T	0.0	¥ 7	47	0.0	¥7	₹
Vos et al 1994 ³⁵	Z	ž	Ϋ́	Z	ž	Ϋ́Α	16.0	2.2‡	0.4 (0.1 to 1.2)†
Moderate use studies (n=8)									
Bowen <i>et al</i> 1984 ⁴⁸	Z	ž	¥Z	ΥZ	Ϋ́Z	Ϋ́Z	3.8§	0.8§	4.9 (1.9 to 12.6)†
Chmielewska-Badora 199836	38.6	9	9.9 (2.6 to 36.8)†	0.4	0.0	NC	ΝΑ	ΥZ	¥Z
	28.1		6.1 (1.8 to 20.4)†						
Fahrer <i>et al</i> 1998 ⁴⁰	ZZ	Z	Y	ž	Z	Ϋ́	0.8	0.8¶	1.0
Fahrer <i>et al</i> 1991 ⁴¹	26.1	3.9	8.6 (2.1 to 35.8)†	1.9-3.1**	Z	Ϋ́Z	0.8	ž	Ϋ́Z
		0.9	5.5 (1.7 to 17.9)†						
Gustafson et al 1993³9	٥	_	9.3 (3.6 to 24.2)†	6.088	1.388	4.8 (1.1 to 20.6)†	Ϋ́	₹Z	Ϋ́Z
		6	0.9 (0.5 to 1.8)†						
Hauser <i>et al</i> 1998 ²³	41.0 to 44.8††	8##	8.4 (4.4 to 15.9)†	ž	Z,	Ϋ́Z	ΥZ	۲	Ϋ́Α
Kuiper <i>et al</i> 1991 ³²	19.7	6.3	3.7 (1.5 to 9.7)	0.9	ž	Ϋ́Z	¥	Ϋ́Z	Ϋ́Z
7hiong et al 1997 ²²	15.2	3.2	5 4 (0 7 to 40 7)+	3.388	a Z	ΔZ	ΔN	∀ Z	▼ Z

*NR, not reported; NA, not applicable; NC, not calculable; OR, odds ratio; RR, relative risk; †calculated with Intercooled Stata, version 6; ‡Data represents reported seroconversion. During the study, one case of Lyme disease was 1.9% for definite disease and reported in the comparison group; **the prevalence of symptomatic disease was 1.9% for definite disease and 3.1% for probable disease; ††1gG and IgM activity to four different antigens were used, range reported is IgG only; ‡‡control data using IgG reactivity only, derived from Hauser et al 1998, table 123; §§lifetime prevalence; for Zhioua et al, 22 the prevalence was for probable disease only.

6.0%. Many of these studies reported lifetime prevalence—that is, diagnosis of disease compatible with Lyme disease at any time in the past. However, only two (Chmielewska-Badora³⁶ and Gustafson *et al*³⁹) reported prevalence of symptomatic Lyme disease among the occupational group under study and a comparison group. Chmielewska-Badora³⁹ documented one case of Lyme disease among 261 farmers and foresters compared with no cases in 50 blood donor controls (odds ratio (OR) not calculable). Gustafson *et al*³⁹ documented a lifetime history of 17 definite and five probable cases of Lyme disease among 362 orienteers compared with two cases of Lyme disease among 150 controls (OR=4.8, 95% confidence interval (95% CI) 1.1 to 20.0). The conclusion of definite disease was based on a past diagnosis of *erythema migrans* made by a physician.

The highest quality epidemiological measure of effect in this review is that of incidence of symptomatic, clinically confirmed disease. Of the 10 studies with high and moderate use, five reported annual incidence of symptomatic, clinically confirmed Lyme disease in the occupational groups under study. Vos et al35 reported both the clinical incidence and seroconversion rates. Overall, relatively few new cases of occupationally acquired Lyme disease were documented in these incidence studies. Both Kuiper et al31 and Vos et al35 found no new cases of symptomatic Lyme disease during their follow up period. Vos et al³⁵ found a risk of seroconversion among the occupational group that was lower than that of the comparison group, although the relative risk did not reach significance. Fahrer et *al*⁴⁰ 41 documented 15 new cases of Lyme disease, equivalent to an annual incidence of 0.8% over the mean follow up interval of 6.5 years. This annual incidence was no different than that found among the non-exposed comparison group. Only Bowen et al48 reported an increased 2 year cumulative incidence of clinical Lyme disease in an outdoor occupational group (3.8%), compared with an indoor occupational group (0.8%), with a relative risk (95% CI) of 4.9 (1.9 to 12.6). These cases were collected retrospectively from among cases reported to the state of New Jersey in 1981 and 1982; the cases that worked at the Naval Weapons Station in Monmouth County were then categorised by indoor and outdoor work.

An important and consistent finding among the studies was that most subjects found to be seropositive, in either prevalence or incidence studies, had no current or past symptoms compatible with Lyme disease. For example, among the studies with high and moderate use, Kuiper *et al*,³¹⁻³² Chmielewska-Badora,³⁶ Hauser *et al*,²³ and Zhioua *et al*²² reported that 82%, 83%, 99%, 100%, and 100% of seropositive subjects were asymptomatic, respectively. Similarly, Vos *et al*,³⁵ and Fahrer *et al*,⁴⁰ reported that 93% and 98% of people with incident seroconversion had no symptoms consistent with Lyme disease.

DISCUSSION

The goal of this exercise was to find if outdoor workers who live, pursue leisure activities, and work in areas endemic for Lyme disease have an increased risk of the disease compared with people who only live and pursue leisure activities in those same areas. Although Lyme disease would seem to be an obvious risk of outdoor work, the scientific literature has not clearly documented the magnitude of the risk of symptomatic, clinically confirmed disease. Many studies have documented the occurrence of Lyme disease in outdoor workers, but few have attempted to document the risk of confirmed, symptomatic disease comparing occupationally exposed people and controls using the current standards of diagnosis of the disease. Most published studies with primary occupational data were cross sectional in design and documented an increased risk of seropositivity among workers compared with controls. However, the use of these studies in guiding decisions about the need to vaccinate outdoor workers in the United

States is limited by: (a) lack of clinical evaluation of subjects; (b) reliance on serological evaluation with ELISA or IFA only, without confirmation by western blot⁵⁸; (c) inadequate assessment of occupational exposure to ticks—for example, lack of measurement of the number of outdoor hours worked or years of service, lack of assessment of protective behaviour or clothing, and failure to adjust for non-occupational exposures; and (d) a predominance of data from European studies, where Borrelia burgdorferi strain differences may alter the clinical presentation of Lyme disease.

In this evaluation of the scientific literature on the occupational risk of Lyme disease, three epidemiological effect measures were evaluated, in order from lowest to highest inferential value: (*a*) odds ratios for seropositivity from cross sectional studies; (*b*) odds ratios for symptomatic, clinically confirmed Lyme disease from cross sectional studies; and (*c*) incidence ratios (relative risks) for symptomatic, clinically confirmed Lyme disease, from longitudinal studies.

Despite the apparent increased risk of seropositivity in occupationally exposed people compared with controls, it should be noted that the choice of an appropriate comparison group can influence the magnitude of this risk. Of the seven studies in table 5 that had an increased risk of seropositivity in the occupational groups under study, five used a comparison group with a decreased residential risk of Lyme disease. Comparing seroprevalence among occupationally exposed people, who generally live in areas endemic for Lyme disease, with controls, who live in areas where Lyme disease is lowest or absent, would bias estimations of occupational risk in the direction of increased risk. Thus, the increased risks of seropositivity for outdoor work presented in table 5 may over estimate the occupational risk.

One consistent deficiency is that less than half of the 41 published studies required clinical confirmation by a physician. What is perhaps most surprising about the published studies is that three studies published in the 1990s all document no increased incidence of symptomatic, clinically confirmed Lyme disease in outdoor workers (Kuiper et al³¹; Vos et al35; and Fahrer et al40). Although one of these studies,31 did not report data from a comparison group, no cases of Lyme disease were found in the occupational group under study. Of the 41 studies, only Bowen et al48 documented an increased incidence of symptomatic Lyme disease among a very specific group of outdoor workers and controls. However, all cases were diagnosed during 1981 and 1982, and thus may not represent either current standards for diagnosis of Lyme disease or current levels of risk after 20 years of experience with the disease among outdoor working populations. Thus, although we think that the findings of Bowen et al⁴⁸ are internally valid, they may not be generalisable, especially to current practice.

Among cross sectional studies that evaluated symptomatic Lyme disease, only one (Gustafson et al39) reported the second most useful effect measure (OR for symptomatic disease); this study reported an increased relative odds for symptomatic Lyme disease over a lifetime, but the methods may raise concerns about recall bias. Many studies documented an increased risk of seropositivity, but most seropositive subjects were asymptomatic, and seropositivity was defined with ELISA or IFA only, which is not the current standard for serological testing. An important point is that, to our knowledge, there are no studies that suggest that asymptomatic seropositive subjects are at risk of developing symptomatic disease or late sequelae of infection. On the contrary, at least one study has reported that there is no increased risk for the development of symptomatic disease in such subjects over an average follow up interval of 6.5 years (Fahrer et al⁴⁰). Serological studies also document significant seroreversion rates; asymptomatic seropositive subjects are seronegative, and still asymptomatic, on repeat testing (Fahrer et al⁴⁰; Schwartz et al54). For example, Schwartz et al54 reported annual seroreversion rates of 43%, 23%, and 53% from 1988-89, 1989-90, and 1990-91, respectively.

Assessment of occupational risk was further limited by the paucity of studies that attempted to define occupational risk factors more carefully (in terms of hours outdoors in specific tick infested habitats, specific high risk tasks, or controlling for use of preventive behaviour) or that controlled for non-occupational risk factors in the assessment of occupational risk. Hours of recreation outdoors, deer sightings near the home, pet ownership, rural residence, and personal preventive behaviour have all been shown to be risk or protective factors for Lyme disease or seropositivity for antibodies to *Borrelia burgdorferi*. 9 10 49 54

Despite the limitations of the scientific literature, it would seem obvious that people who work outdoors in tick infested areas should be at increased risk of Lyme disease; however, an increased risk of symptomatic, clinically confirmed Lyme disease has not been documented in outdoor workers. Many of the studies were not specifically designed to evaluate the occupational risk of symptomatic, clinically confirmed Lyme disease, and the current serological testing guidelines were developed after most of these studies were published. However, the three best published epidemiological studies do not suggest that an occupational risk exists (Vos *et al*⁷⁵; Kuiper *et al*⁷¹; Fahrer *et al*⁷⁰). It may be that workers in tick infested habitat become knowledgeable about the disease and use personal preventive behaviour to minimise their risk. ^{9 55}

Several studies suggest that such personal protective behaviour as tick checks, tucking trousers into socks, or use of permethrin or DEET may decrease the risk of seropositivity or tick bites. Antitick saliva antibody (ATSA) and antirecombinant tick calreticulin antibody are two biomarkers of tick exposure that have been used in epidemiological studies of exposure to ticks.^{8 59 60} For example, in a study of military personnel on manoeuvres in tick infested areas of Arkansas, people who tucked their trousers into their socks were significantly less likely to be ATSA seropositive than subjects who did not tuck in their trousers (OR 2.8 (95% CI 1.1 to 7.1, Schwartz 1996).⁸ Similarly, outdoor workers in New Jersey who did not use insect repellants were more likely to be ATSA seropositive (OR (95% CI) = 2.0 (1.0-4.0), Schwartz and Goldstein).⁹

Environmental application of insecticides has been shown to decrease the abundance of ticks, but this is more likely to be useful in the prevention of residentially acquired Lyme disease than in the prevention of occupational disease. If It is also known that duration of tick feeding is an important determinant of the risk of infection, so tick checks and early tick removal are likely to be effective in the prevention of the disease. It should be noted, however, that other studies have not shown that personal preventive behaviour is effective in disease prevention. All of these strategies have been recommended by the Centers for Disease Control as effective measures to decrease the risk of acquiring Lyme disease.

There are no other personal strategies to prevent Lyme disease that have been proved to prevent the disease as effectively as the Lyme disease vaccine. The Lyme vaccine efficacy study was a randomised, placebo controlled trial in 10 936 subjects aged 15–70 years. ⁶² Lyme disease was carefully documented with culture, polymerase chain reaction, or western blot sero-conversion among subjects with symptoms compatible with Lyme disease. Among study subjects under the age of 65, after three doses the vaccine was 90% effective in preventing laboratory confirmed, symptomatic Lyme disease. The Centers for Disease Control, The Medical Letter, and other authors have all commented on use of the vaccine. ^{1 13 14}

Another important factor to consider is the natural history of Lyme disease compared with other diseases preventable by vaccine. Lyme disease is non-fatal, relatively easy to diagnose, and relatively easy to treat with oral antibiotics. By contrast, hepatitis B virus, for example, can be fatal, can have a chronic carrier state, and is not effectively treated. Although vaccination against Lyme disease may help to decrease morbidity, it would seem less imperative than for an organism such as hepatitis B virus.

When deciding whether or not to vaccinate a workforce against Lyme disease, the occupational physician should consider several factors—the strength of the scientific evidence presented here, the availability of resources, the history of risk of Lyme disease in the workforce under consideration, and the concerns of workers. The scientific literature shows an increased risk of occupational exposure to Borrelia burgdorferi, as assessed by the seroprevalence studies, but fails to document an increased risk of development of symptomatic Lyme disease. The lack of documented clinical risk makes it difficult to rely exclusively on the current scientific literature. Factors that may favour vaccination would include documented increased risk of Lyme disease in the workforce under consideration for vaccination; a high level of concern about the disease from workers; high level of outdoor activity in tick infested areas; poor compliance with personal protective clothing or behaviour; and availability of financial resources for occupational health programmes. Further studies that explicitly assess the risk of developing occupationally acquired Lyme disease are necessary to help guide the physician on whether or not to vaccinate.

ACKNOWLEDGEMENTS

This research was supported in part by SmithKline Beecham Pharmaceuticals.

Authors' affiliations

- J D Piacentino, B S Schwartz, Division of Occupational and Environmental Health, Department of Environmental Health Sciences, Johns Hopkins Bloomberg School of Hygiene and Public Health, Baltimore, MD, USA
- J D Piacentino, B S Schwartz, Department of Medicine, Johns Hopkins Bloomberg School of Medicine
- B S Schwartz, Department of Epidemiology, Johns Hopkins Bloomberg School of Hygiene and Public Health

REFERENCES

- 1 Centers for Disease Control. Recommendations for the use of Lyme disease vaccine, recommendations of the Advisory Committee on Immunization Practices (ACIP). MMWR Morb Mortal Wkly Rep 1999:48:1-17
- 2 Centers for Disease Control. Surveillance for Lyme disease, United States, 1992–1998. MMWR Morb Mortal Wkly Rep 2000;49:1-11.
- 3 Coyle BS, Strickland GT, Liang YY, et al. Public impact of Lyme disease in Maryland. J Infect Dis 1996;173:1260-2.
- 4 Meek JI, Roberts CL, Smith E, et al. Underreporting of Lyme disease by Connecticut physicians, 1992. Journal of Public Health Management and Practice 1996:2:61-5
- 5 Lastavica CC, Wilson ML, Berardi VP, et al. Rapid emergence of a focal epidemic of Lyme disease in coastal Massachusetts. N Engl J Med 1989;**320**:133–7.
- 6 Schwartz B, Goldstein M. Lyme disease: a review for the occupational physician. J Occup Med 1989;31:735–42.
 7 Parrott C, Johnson K, Strauss S, et al. Lyme disease in outdoor workers.
- on Assateague Island: high tick-exposure but low disease risk. Md Med J
- 8 Schwartz BS, Sanchez J, Sanders M, et al. Tick avoidance behaviors associated with a decreased risk of anti-tick salivary gland protein antibody seropositivity in miliatry personnel exposed to Amblyomma Americanum in Arkansas. Am J Trop Med Hyg 1996;44:410–16.
 9 Schwartz BS, Goldstein M. Lyme disease in outdoor workers: risk
- factors, preventive measures, and tick removal methods. Am J Epidemiol 1990;**131**:877-85.
- 10 Shadick N, Daltroy L, Phillips C, et al. Determinants of tick-avoidance behaviors in an endemic area for Lyme disease. Am J Prev Med 1997;13:265-70.
- Curran K, Fish D, Piesman J. Reduction of nymphal Ixodes dammini in a residential suburban landscape by area application of insecticides. J Med Entomol 1993;30:107-13.
- 12 Hayes E, Maupin G, Mount G, et al. Assessing the effectiveness of local Lyme disease control. Journal of Public Health Management and Practice 1999;**5**:86–94.
- 13 Marwick C. Guarded endorsement for Lyme disease vaccine. JAMA 1998;**279**:1937–8.
- 14 Meltzer M, Dennis D, Orloski K. The cost effectiveness of vaccinating against Lyme disease. Emerg Infect Dis 1999;**5**:1–8.
- 15 Steere A, Sikand V, Meurice F, et al. Vaccination against Lyme disease with recombinant Borrelia burgdorferi outer-surface lipoprotein A with adjuvant. N Engl J Med 1998;339:209-15.

- 16 The Medical Letter. Lyme disease vaccine. Med Lett Drugs Therap 1999;41:29–30.
- 17 Stanchi N, Balaque L. Lyme disease: antibodies against Borrelia burgdorferi in farm workers in Argentina. Rev Saude Publica 1993:**27**:305-7
- 18 Schmutzhard E, Stanek G, Pletschette M, et al. Infections following tickbites. Tick-borne encephalitis and Lyme Borreliosis. Infection 1988;**16**:269–72.
- 19 Golubic D, Rijpkema S, Tkalec-Makovec N, et al. Epidemiologic, ecologic and clinical characteristics of LB in NW Croatia. Acta Med Croatica 1998;**52**:7–13.
- 20 Oksi J Viljanen M. Tick bites, clinical symptoms of Lyme borreliosis, and borrelia antibody responses in Finnish army recruits training in an endemic region during summer. *Mil Med* 1995;**160**:453–6.
- Christiann F, Rayet P, Patey O, et al. Epidemiology of Lyme disease in France: Lyme borreliosis in the region of Berry Sud. Eur J Epidemiol 1996:**12**:479-83.
- 22 **Zhioua E**, Rodhain F, Binet PH, *et al.* Prevalence of antibodies to Borrelia burgdorferi in forestry workers of Ile de France, France. Eur J Epidemiol 1997;**13**:959–62
- 23 Hauser U, Krahl H, Peters H, et al. Impact of strain heterogeneity on
- Lyme disease serology in Europe: comparison of EUSA using different species of Borrelia burgdorferi sl. J Clin Microbiol 1998;36:427–36.
 Rath P, Ibershoff B, Mohnhaupt A, et al. Seroprevalence of Lyme borreliosis in forestry workers from Brandenburg, Germany. Eur J Clin Microbiol Infect Dis 1996;15:372–7.
- 25 Robertson J, Gray J, MacDonald S, et al. Seroprevalence of Borrelia burgdorferi sensu lato infection in blood donors and park rangers in relation to local habitat. Zentralb Bakteriol 1998;288:293–301.
- Nuti M, Amaddeo D, Crovatto M, et al. Infections in an alpine environment. Am J Trop Med Hyg 1993;48:20–5.
 Santino I, Iori A, Sessa R, et al. Borrelia burgdorferi and Ehrlichia
- chaffeensis in the National Park of Abruzzo. FEMS Microbiol Lett 1998;164:1-6.
- 28 Ikushima M, Yamada F, Kawahashi S, et al. Antibody response to OspC-I synthetic peptide derived from outer surface protein C of Borrleia burgdorferi in sera from Japanese forestry workers. *Epidemiol Infect* 1999;**122**:429–33.
- 29 Nakama H, Muramatsu K, Uchikawa K, et al. Possibility of Lyme disease as an occupational disease: seroepidemiological study of regional residents. *Asia Pac J Public Health* 1994;**7**:214–7.
- Montejunas L, Bunikis J, Barbour A, et al. Lyme borreliosis in Lithuania. Scand J Infect Dis 1994;26:149–55.
- 31 Kuiper H, de Jongh B, Nauta A, et al. One year follow up study to assess the prevalence and incidence of Lyme borreliosis among Dutch forestry workers. Eur J Clin Microbiol Infect Dis 1993;12:413–8
- 32 Kuiper H, Dam A, van Charante A, et al. Lyme borreliosis in Dutch forestry workers. J Infect 1991;23:279-86.
- 33 Van Charante M, Groen J, Mulder P, et al. Occupational risks of zoonotic infections in Dutch forestry workers and muskrat catchers. Eur J Epidemiol 1998;14:109-16.
- 34 Van Charante M, Groen J, Osterhaus A. Risk of infections transmitted by arthropods and rodents in forestry workers. Eur J Epidemiol 1994;**10**:349-51.
- 35 Vos K, Van Dam A, Kuiper H, et al. Seroconversion for Lyme borreliosis 35 Vos K, Van Dam A, Nuipei Fi, et al. Seroconversion for Lynne Bottonios among Dutch military. Scand J Infect Dis 1994;26:427–34.
 36 Chmielewska-Badora J. Seroepidemiologic study on LB in the Lublin
- region. Ann Agric Environ Med 1998;5:183-6.
- 37 Arteaga F, Golightly M, Perez A, et al. Disparity between serological reactivity to B burgdorferi and evidence of past disease in a high risk group. Clin Infect Dis 1998;**27**:1210–3.
- 38 Oteo J, Artola M, Casas J, et al. Epidemiology and prevalence of seropositivity against Borrelia burgdorferi antigen in La Rioja, Spain. Rev Epidemiol Sante Publique 1992;40:85–92.
- 39 Gustafson R, Forsgren M, Gardulf A, et al. Antibody prevalence and clinical manifestations of Lyme borreliosis and tick-borne encephalitis in Swedish orienteers. Scand J Infect Dis 1993;25:605-11.
- 40 **Fahrer H**, Sauvain S, Zhioua E, *et al*. Long term survey in a population at risk for LB; what happens to the seropositive individuals? Eur J Epidemiol 1998;**14**:117-23.
- 41 Fahrer H, Linden S, Sauvain M, et al. The prevalence and incidence of clinical and asymptomatic Lyme borreliosis in a population at risk. J Infect Dis 1991;163:305-10.
- 42 Baird A, Gilliew J, Bone F, et al. Prevalence of antibody indicating Lyme disease in farmers in Wigtownshire. *BMJ* 1989;**29**9:836–7.

 43 **Gregory RP**, Green A, Merry R. Lyme disease in military personnel. *J R*
- Army Med Corps 1993;139:11-13.
- 44 Guy E, Martyn C, Bateman D, et al. Lyme disease prevalence and clinical importance of Borrelia burgdorferi specific IgG in forestry
- workers. Lancet 1989;i:484–5.

 45 Morgan P, Cutler S, Wright D. Borrelia burgdorferi infection in UK workers at risk of tick bites. Lancet 1989;i:789–90.
- 46 **Reese D**, Axford J. Evidence for LD in urban park workers: a potential new health hazard for city inhabitants. British Journal of Rheumatology 1994;33:123-8.
- 47 Thomas D, Sillis M, Coleman T, et al. Low rates of erlichiosis and Lyme
- Hornds J., Sillis M., Colental T., et al. Low Tales of entitions and sylle borreliosis in English farmworkers. Epidemiol Infect 1998;121:609–14.
 Bowen G., Schulze T., Hayne C., et al. A focus of Lyme disease in Monmouth County, New Jersey. Am J Epidemiol 1984;120:387–94.
 Goldstein M., Schwartz B., Friedmann C., et al. Lyme disease in New
- Jersey outdoor workers: a statewide survey of seroprevalence and tick exposure. Am J Public Health 1990;80:1225-9.

- 50 **Klein J.** Lyme disease in unwary physicians. *JAMA* 1995;**274**:383–4. 51 **Lane R**, Manweiler S, Stubbs H, *et al.* Risk factors for Lyme disease in a small rural community in northern California. Am J Epidemiol 1992;**136**:1358-68.
- 52 Ley C, Olshen E, Reingold A. Case-control study of risk factors for incident Lyme disease in California. Am J Epidemiol 1995; **142**:S39–47 53 **Schwartz BS**, Goldstein M, Childs J. Antibodies to Borrelia burgdorferi
- and tick salivary gland proteins in NJ outdoor workers. Am J Public Health 1993;83:1746–8.
- 54 **Schwartz BS**, Goldstein M, Childs J. Longitudinal study of Borrelia burgdorferi infection in NJ outdoor workers. Am J Epidemiol
- 55 Smith P, Benach J, White D, et al. Occupational risk of Lyme disease in endemic areas of NY state. Ann NY Acad Sci 1988;539:289–301.
 56 Zhioua E, Gern L, Aeschlimann A, et al. Longitudinal study of Lyme
- borreliosis in a high risk population in Switzerland. Parasite 1998:**5**:383-6
- 57 Christiann F, Rayet P, Patey O, et al. Lyme borreliosis in central France: a sero-epidemiologic examination involving hunters. Eur J Epidemiol 1997:13:885.

- 58 Centers for Disease Control. Recommendations for test performance and interpretation from the 2nd National Conference on Serologic Diagnosis of Lyme Disease. MMWR Morb Mortal Wkly Rep 1995;**44**:590–1.
- 59 Sanders ML, Jaworski D, Sanchez JL, et al. Antibody to a cDNA-derived calreticulin protein from Amblyomma americanum is a biomarker of tick exposure in humans. Am J Trop Med Hyg 1998;**59**:279–85. 60 **Schwartz BS**, Ribeiro JMC, Goldstein MD. Anti-tick antibodies: an
- epidemiologic tool in Lyme disease research. *Am J Epidemiol* 1990;**132**:58–66.
- 61 Piesman J, Mather T, Sinsky RJ, et al. Duration of tick attachment and B. burgdorferi transmission. J Ćlin Microbiol 1987;25:557-8
- Steere A, Sikand V, Meurice F, et al. Prevalence of antibody indicating Lyme disease in farmers in Wigtownshire. BMJ 1989;299:836–7.
 The Occupational Safety and Health Act of 1970. P.L. 91–596, 84 Stat. 1590, 29 U.S.C.A. §§651–78, as amended by P.L. 95–239,
- 64 Mast EE. Prevention of hepatitis B virus infection in healthcare workers. In: Ellis RW, ed. Hepatitis B vaccines in clinical practice. New York: Marcel Dekker, 1993;295-307.

Answers to multiple choice questions on Complementary and alternative medicine: what is it all about? by E Ernst and A Fugh-Berman on pages 140-144

- (1)(a) true;
 - (b) false—few therapies are whole systems, many are discrete treatments;
 - (c) false—it is "diagnosis, treatment and/or prevention which complements mainstream medicine by contributing to a common whole, by satisfying a demand not met by orthodoxy or by diversifying the conceptual frameworks of medicine";
 - (d) true
- (2)(a) true;

 - false—complementary therapist visits were more numerous by 70%;
 - (d) false: in the USA, 72% did not tell their physician
- (3)(a) false—there is no evidence in support of this hypothesis;
 - (b) true;
 - (c) true—at least in breast cancer patients;
 - (d) true
- (4)(a) false—the figure is 16%;
 - (b) true;
 - (c) false—there is clear evidence that the same evidence is required for complementary medicine as is required for conventional medicine;
 - (d) false—evidence exists
- (5)(a) false—there is limited evidence that it is more effective;
 - (b) false—the evidence suggests the effects of homoeopathy are not completely due to placebo;
 - false—the evidence is not convincing;
 - (d) true